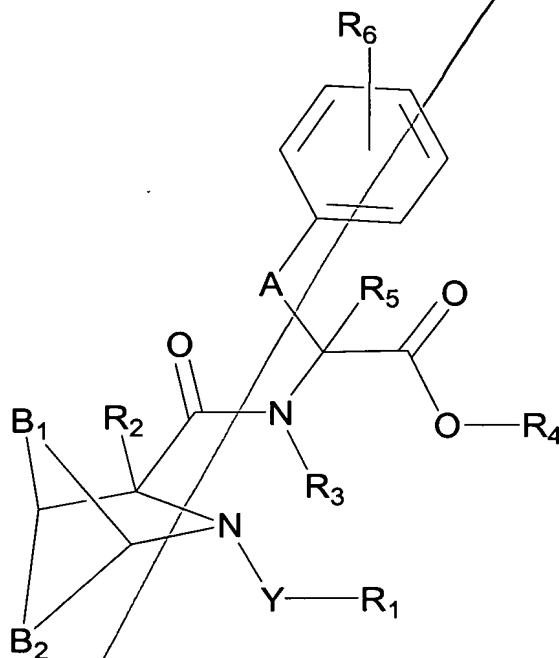


WHAT IS CLAIMED IS:

1. A compound of Formula (I):



Formula (I)

wherein

Y is selected from the group consisting of a bond, -C(O)-, -C(O)O-, -C(O)NH- and -SO₂-;

R₁ is selected from the group consisting of R₇ and R₈;

R₂, R₃, R₄ and R₅ are independently selected from the group consisting of a bond, hydrogen and C₁₋₈alkyl; wherein C₁₋₈alkyl is optionally substituted with one to three substituents independently selected from R₉, provided that R₂, R₃, R₄ or R₅ can only be a bond when forming a monocyclic ring wherein the following monocyclic rings may be formed from R₂, R₃, R₄ and R₅;

when R₂ and R₃ comprise a bond and C₁₋₈alkyl or optionally when both R₂ and R₃ are C₁₋₈alkyl, R₂ and R₃ together with the atoms to which each is attached will form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

and C₂₋₈alkenylene optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C₁₋₈)alkyl, hydroxy(C₁₋₈)alkoxy, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃;

and pharmaceutically acceptable salts, racemic mixtures, diastereomers and enantiomers thereof.

10

2. The compound of claim 1 wherein Y is selected from the group consisting of -C(O)- and -SO₂-.

3. The compound of claim 1 wherein Y is selected from -SO₂-.

4. The compound of claim 1 wherein R₁ is selected from R₇.

5. The compound of claim 1 wherein R₂, R₃, R₄ and R₅ are independently selected from the group consisting of hydrogen and C₁₋₄alkyl.

6. The compound of claim 1 wherein R₂, R₃, R₄ and R₅ are independently selected from the group consisting of hydrogen and methyl.

7. The compound of claim 1 wherein R₆ is optionally present and is one to three substituents independently selected from the group consisting of halogen, C₁₋₈alkoxy, R₁₀, R₁₂, -N(R₁₁)C(O)-R₁₀, -N(R₁₁)C(O)-R₁₂, -N(R₁₁)SO₂-R₁₀, -N(R₁₁)C(O)-N(R₁₁,R₁₂), -N(R₁₁)C(O)-N(R₁₂,R₁₇), -OC(O)-N(R₁₁,R₁₂), -OC(O)-N(R₁₂,R₁₇), -OC(O)-R₁₀ and R₁₀-(C₁₋₈)alkoxy.

8. The compound of claim 1 wherein R₆ is optionally present and is one to three substituents independently selected from the group consisting of halogen, C₁₋₄alkoxy, R₁₀, R₁₂, -N(R₁₁)C(O)-R₁₀, -N(R₁₁)C(O)-R₁₂, -N(R₁₁)SO₂-R₁₀, -N(R₁₁)C(O)-N(R₁₁,R₁₂), -N(R₁₁)C(O)-N(R₁₂,R₁₇),

-OC(O)-N(R₁₁,R₁₂), -OC(O)-N(R₁₂,R₁₇), -OC(O)-R₁₀ and R₁₀-(C₁₋₄)alkoxy.

9. The compound of claim 1 wherein R₆ is optionally present and is one to two substituents independently selected from the group consisting of R₁₀, -N(R₁₁)C(O)-R₁₀, -N(R₁₁)C(O)-N(R₁₁,R₁₂), -N(R₁₁)C(O)-N(R₁₂,R₁₇), -OC(O)-N(R₁₁,R₁₂), -OC(O)-N(R₁₂,R₁₇) and R₁₀-methoxy.
10. The compound of claim 1 wherein R₇ is selected from the group consisting of aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, C₁₋₈alkylcarbonyl, C₁₋₈alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃; and, wherein the aryl and heteroaryl substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃.
11. The compound of claim 1 wherein R₁₀ is selected from the group consisting of cycloalkyl, heterocyclyl, aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₁₋₈alkoxy, C₁₋₈alkoxycarbonyl, carboxyl, arylcarbonyl, arylsulfonyl, -CF₃ and -OCF₃; wherein cycloalkyl and heterocyclyl are optionally substituted with one to three oxo substituents ; and, wherein the aryl portion of the arylcarbonyl substituent is optionally substituted with one to five substituents independently selected from C₁₋₈alkoxy.
12. The compound of claim 1 wherein R₁₀ is selected from the group consisting of cyclopropyl, 1,3-dihydro-2H-isoindolyl, 2-azabicyclo[2.2.2]octyl, piperidinyl, morpholinyl, phenyl, naphthalenyl,

thienyl, 1*H*-pyrrolyl and pyridinyl; wherein cyclopropyl, piperidinyl, morpholinyl, phenyl, naphthalenyl, thienyl, 1*H*-pyrrolyl and pyridinyl are optionally substituted with one to four substituents independently selected from the group consisting of chlorine, fluorine, bromine, methyl, isopropyl, *t*-butyl, methoxy, *t*-butoxycarbonyl, carboxyl, phenylcarbonyl, -CF₃ and -OCF₃; wherein 1,3-dihydro-2*H*-isoindolyl is optionally substituted with oxo; wherein 2-azabicyclo[2.2.2]octyl is optionally substituted with phenylsulfonyl, and, wherein the phenyl portion of the phenylcarbonyl substituent is optionally substituted with one to two substituents independently selected from methoxy.

13. The compound of claim 1 wherein R₁₂ is selected from the group consisting of C₁₋₈alkyl and C₂₋₈alkynyl optionally substituted on a terminal carbon with R₁₄.
14. The compound of claim 1 wherein R₁₂ is selected from the group consisting of C₁₋₄alkyl and C₂₋₄alkynyl optionally substituted on a terminal carbon with R₁₄.
15. The compound of claim 1 wherein R₁₂ is selected from the group consisting of *t*-butyl and ethynyl; wherein ethynyl is optionally substituted on a terminal carbon with a substituent independently selected from R₁₄.
16. The compound of claim 1 wherein R₁₄ is selected from the group consisting of aryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, C₁₋₈alkylcarbonyl, C₁₋₈alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, *N*-(C₁₋₈alkyl)amino, *N,N*-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃; and, wherein the aryl and heteroaryl substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino,

$N-(C_{1-8}\text{alkyl})\text{amino}$, $N,N-(C_{1-8}\text{dialkyl})\text{amino}$, $-\text{CF}_3$ and $-\text{OCF}_3$.

17. The compound of claim 1 wherein R_{11} is selected from the group consisting of hydrogen and $C_{1-4}\text{alkyl}$.

18. The compound of claim 1 wherein R_{11} is hydrogen.

19. The compound of claim 1 wherein A is selected from the group consisting of methylene and ethylene.

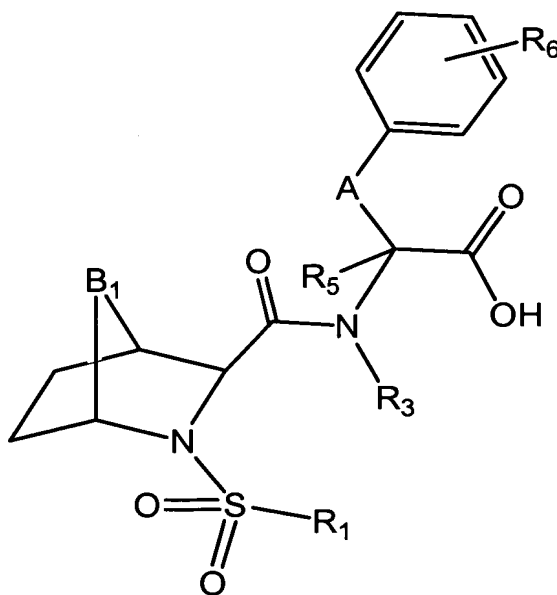
20. The compound of claim 1 wherein B_1 and B_2 are independently selected from the group consisting of $C_{1-4}\text{alkylene}$ and $C_{2-4}\text{alkenylene}$ optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C_{1-4})alkyl, hydroxy(C_{1-4})alkoxy, $C_{1-4}\text{alkyl}$, $C_{2-4}\text{alkenyl}$, $C_{2-4}\text{alkynyl}$, $C_{1-4}\text{alkoxy}$, carboxyl, amino, $N-(C_{1-4}\text{alkyl})\text{amino}$, $N,N-(C_{1-4}\text{dialkyl})\text{amino}$, $-\text{CF}_3$ and $-\text{OCF}_3$.

20 21. The compound of claim 1 wherein B_1 and B_2 are independently selected from the group consisting of $-\text{CH}_2-$, $-(\text{CH}_2)_2-$ and $-(\text{CH})_2-$ optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C_{1-4})alkyl, hydroxy(C_{1-4})alkoxy, $C_{1-4}\text{alkyl}$, $C_{2-4}\text{alkenyl}$, $C_{2-4}\text{alkynyl}$, $C_{1-4}\text{alkoxy}$, carboxyl, amino, $N-(C_{1-4}\text{alkyl})\text{amino}$, $N,N-(C_{1-4}\text{dialkyl})\text{amino}$, $-\text{CF}_3$ and $-\text{OCF}_3$.

21 22. The compound of claim 1 wherein B_1 is selected from the group consisting of $-\text{CH}_2-$, $-(\text{CH}_2)_2-$ and $-(\text{CH})_2-$ optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C_{1-4})alkyl, hydroxy(C_{1-4})alkoxy, $C_{1-4}\text{alkyl}$, $C_{2-4}\text{alkenyl}$, $C_{2-4}\text{alkynyl}$, $C_{1-4}\text{alkoxy}$, carboxyl, amino, $N-(C_{1-4}\text{alkyl})\text{amino}$, $N,N-(C_{1-4}\text{dialkyl})\text{amino}$, $-\text{CF}_3$ and $-\text{OCF}_3$; and, wherein, B_2 is selected from $-(\text{CH}_2)_2-$.

- 22/23. The compound of claim 1 wherein B_1 is selected from the group consisting of $-\text{CH}_2-$, $-(\text{CH}_2)_2-$ and $-(\text{CH})_2-$.

- 5 23/24. The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:



wherein B_1 , R_1 , R_3 , R_5 , A and R_6 are dependently selected from the group consisting of:

B_1	R_1	R_3	R_5	A	R_6
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-(2,6- Cl_2)Ph;
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-(2,4,6- Cl_3)Ph;
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-[2,6-(OMe) $_2$]Ph;
CH_2	Ph	H	H	CH_2	4-NHC(O)-(2,6- F_2)Ph;
$(\text{CH}_2)_2$	Ph	H	H	CH_2	4-NHC(O)-(2,6- Cl_2)Ph;
$(\text{CH}_2)_2$	Ph	H	H	CH_2	4-[2,6-(OMe) $_2$]Ph;
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-(2-Me)Ph;
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-(2-Cl)Ph;
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-(2,6- F_2)Ph;
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-(2- CF_3)Ph;
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-(2- OCF_3)Ph;
$(\text{CH}_2)_2$	4-Tol	H	H	CH_2	4-NHC(O)-(2-Br)Ph;
$(\text{CH}_2)_2$	Ph	H	H	CH_2	4-NHC(O)-(2,6- F_2)Ph;
CH_2	Ph	H	H	CH_2	4-NHC(O)-(2,6- Cl_2)Ph;

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(CH ₂) ₂	4-Tol	H	H	CH ₂	4-[2,6-(OMe) ₂]Ph;
CH ₂	Ph	H	H	CH ₂	4-NHC(O)-[2,6-(OMe) ₂]Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-CC-(4- <i>t</i> -butyl)Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-CC-Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-NHC(O)-Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-NHC(O)-[4-C(O)-[2,5-(OMe) ₂]Ph]Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-NHC(O)-CH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-NH-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-OCH ₂ -Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-NHC(O)-(2,4,6-isopropyl ₃)Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-(1 <i>H</i> -pyrrol-1-yl);
(CH ₂) ₂	4-Tol	H	H	CH ₂	4-Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-NH-(2,6-F ₂)Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	3-NHC(O)-(2,6-F ₂)Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	3-NHC(O)-[2,6-(OMe) ₂]Ph;
(CH ₂) ₂	4-Tol	H	H	CH ₂	3-NHC(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	CH ₃	CH ₂	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	CH ₃	H	CH ₂	4-NHC(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-(2,4,6-F ₃)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-(2,3,5,6-F ₄)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-O- <i>t</i> -butoxy;
(CH ₂) ₂	Ph	H	H	(CH ₂) ₂	---
(CH ₂) ₂	Ph	H	H	CH ₂	4-(1,3-dihydro-1,3-dioxo-2 <i>H</i> -isoindol-2-yl);
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-(2-CO ₂ H)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-(2,5-diMe-1 <i>H</i> -pyrrol-1-yl);
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-4-pyridinyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHSO ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-OC(O)-N(CH ₃) ₂ ;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-(1- <i>t</i> -butoxycarbonyl)4-piperidinyl;
(CH ₂) ₂	4-FPh	H	H	CH ₂	4-NHC(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	4-FPh	H	H	CH ₂	4-NHC(O)-[2,6-(OMe) ₂]Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-OC(O)-4-morpholinyl;

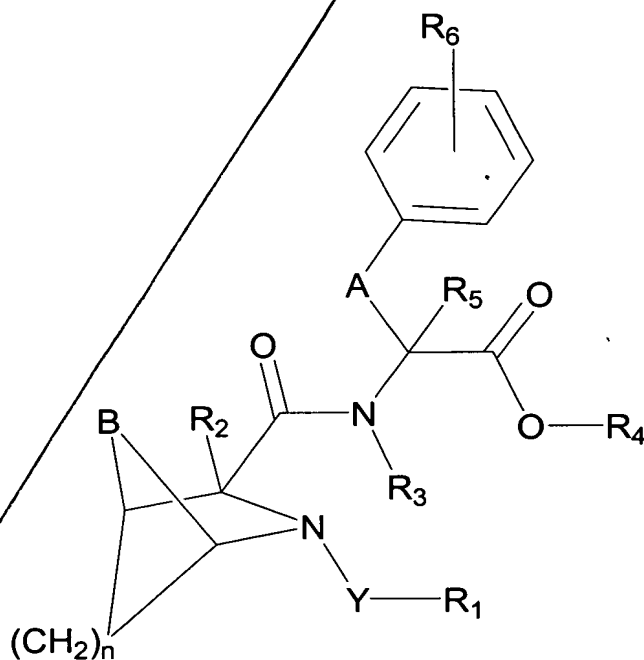
(CH ₂) ₂	Ph	H	H	CH ₂	4-OC(O)N(<i>iso</i> -propyl) ₂ ;
(CH ₂) ₂	Ph	H	H	CH ₂	4- <i>t</i> -butyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-4-piperidinyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-(3,5-Cl ₂)4-pyridinyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-NMe ₂ ;
(CH ₂) ₂	Ph	H	H	CH ₂	3-F-4-[OCH ₂ (2,6-Cl ₂)Ph] ;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-OC(O)-NMe ₂ ;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)- <i>t</i> -butyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-(2-OMe)1-naphthalenyl;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-NHC(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-cyclopropyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-(2,2,3,3-Me ₄)cyclopropyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)- <i>iso</i> -propyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-(2-SO ₂ Ph)-2-azabicyclo[2.2.2]oct-3-yl;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-NHC(O)-(3,5-Cl ₂)4-pyridinyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-NHC(O)-(2-Me)cyclopropyl;
(CH ₂) ₂	Ph	H	H	CH ₂	4-(2,6-diMe)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-(2,6-Cl ₂)Ph;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-(2,6-Cl ₂)Ph;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-(2,6-diMe)Ph;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-[2,6-(OMe) ₂]Ph;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-(4-fluoro-1,3-dihydro-1,3-dioxo-2 <i>H</i> -isoindol-2-yl);
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-NHC(O)-NMe ₂ ;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-OC(O)-NMe ₂ ;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-OC(O)-(4-morpholinyl);
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-OC(O)-(4-Me-1-piperazinyl);
(CH ₂) ₂	Ph	H	H	CH ₂	4-OC(O)-(4-Me-1-piperazinyl);
(CH ₂) ₂	Ph	H	H	CH ₂	4-N(Me)C(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-N(Me)C(O)-(3,5-Cl ₂)4-pyridinyl;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-N(Me)C(O)-(3,5-Cl ₂)4-pyridinyl;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-N(Me)C(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-OCH ₂ -(2,6-Cl ₂)Ph;
(CH ₂) ₂	2-Thi	H	H	CH ₂	4-(1,3-dihydro-1,3-dioxo-2 <i>H</i> -isoindol-2-yl);
(CH ₂) ₂	Ph	H	H	CH ₂	4-(1,3-dihydro-4,7-dimethyl-1,3-dioxo-2 <i>H</i> -isoindol-2-yl);

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(CH ₂) ₂	2-Thi	H	H	CH ₂	4-(1,3-dihydro-4,7-dimethyl-1,3-dioxo-2 <i>H</i> -isoindol-2-yl);
CH ₂	2-Thi	H	H	CH ₂	4-NHC(O)-(3,5-Cl ₂)4-pyridinyl;
CH ₂	2-Thi	H	H	CH ₂	4-NHC(O)-(2,6-Cl ₂)Ph;
(CH ₂) ₂	Ph	H	H	CH ₂	4-(1,1-dioxido-3-oxo-1,2-benzisothiazol-2(3 <i>H</i>)-yl);
(CH ₂) ₂	Ph	H	H	CH ₂	4-(4-chloro-1,3-dihydro-1,3-dioxo-2 <i>H</i> -isoindol-2-yl);
and,					
(CH ₂) ₂	Ph	H	H	CH ₂	4-(7,9-dioxo-8-azaspiro[4.5]dec-8-yl);

and pharmaceutically acceptable salts, racemic mixtures, diastereomers and enantiomers thereof.

25. A compound having Formula (II):



Formula (II)

wherein

Y is selected from the group consisting of -C(O)- and -SO₂-;

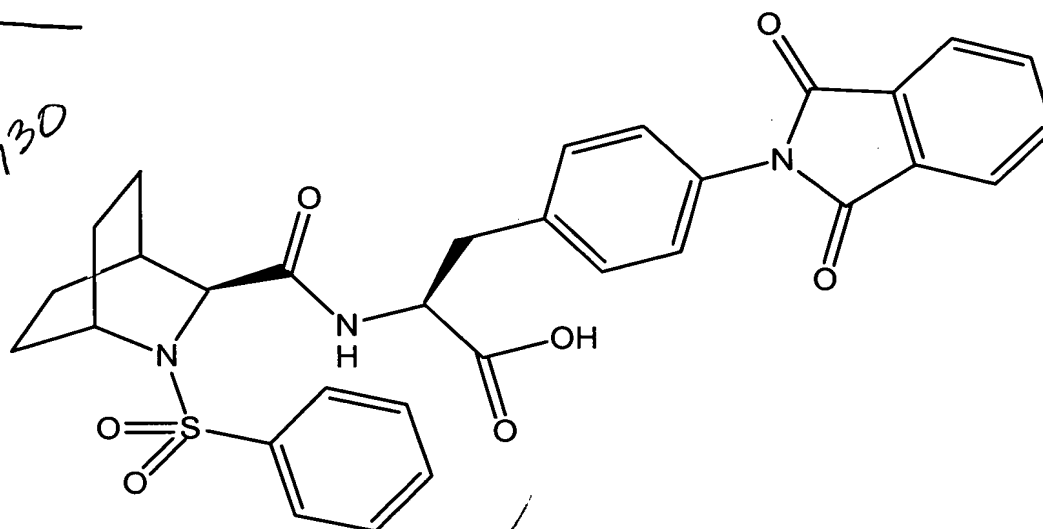
R₁ is selected from the group consisting of R₇ and R₈;

R₂, R₃, R₄ and R₅ are independently selected from the group consisting of a bond, hydrogen and C₁₋₈alkyl; wherein C₁₋₈alkyl is optionally substituted with

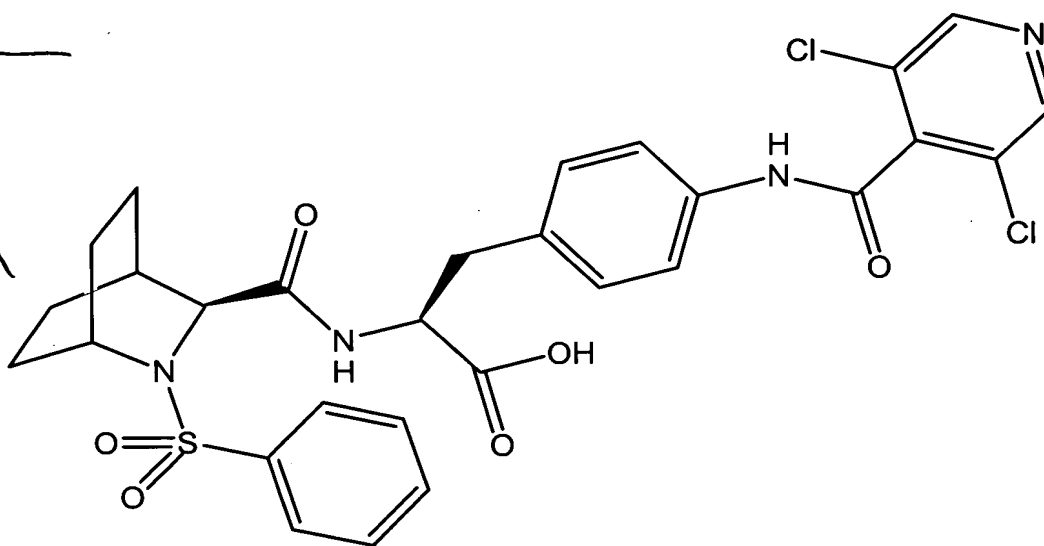
in the presence of appropriate coupling agents, bases and solvents to form the compound of Formula (II).

The process of claim 25 wherein R_{15} is selected from the group consisting of hydroxy, iodine, bromine and NO_2 .

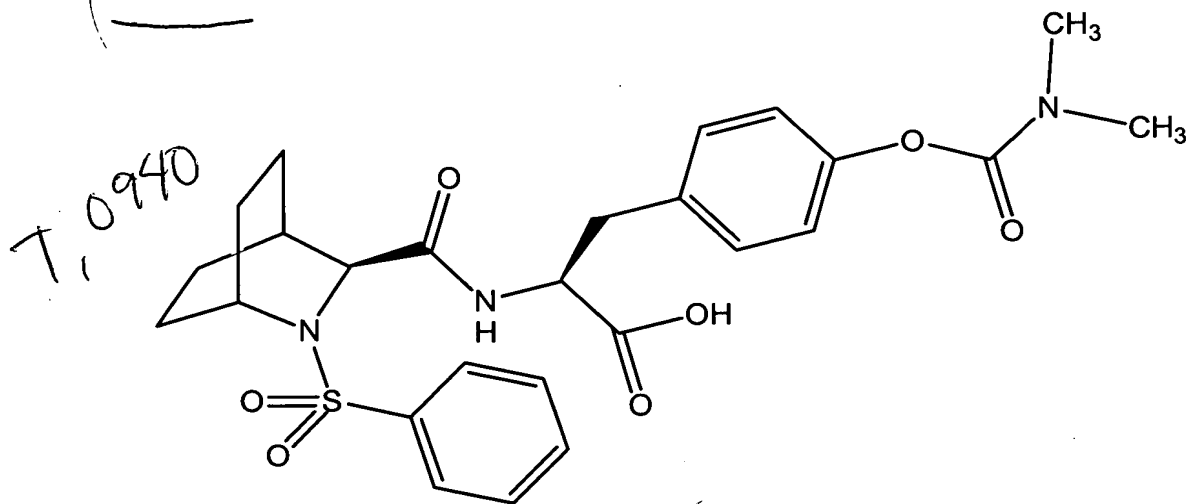
The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:



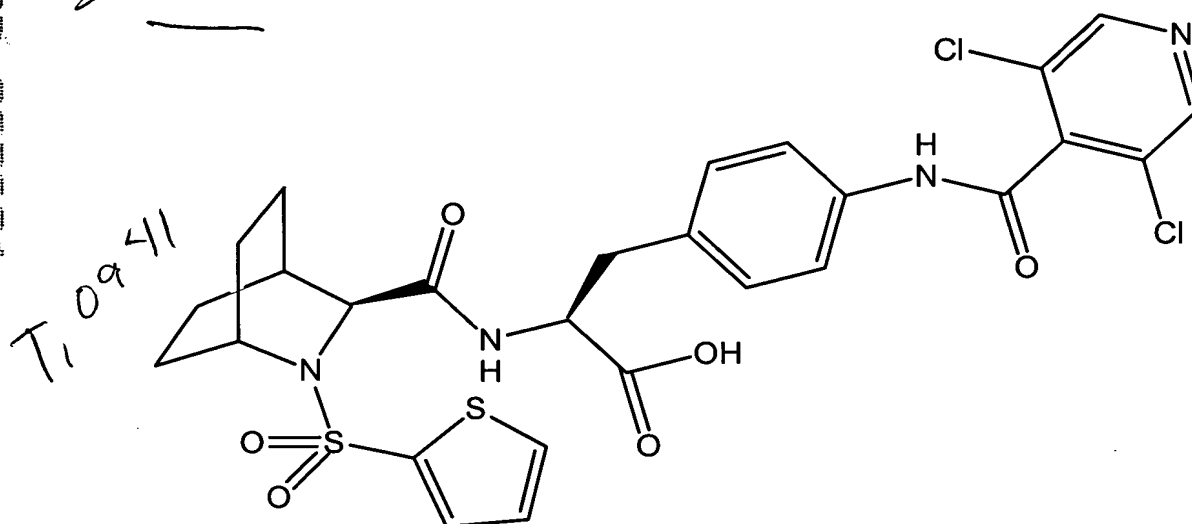
The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:



30. The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:



31. The compound of claim 1 wherein the compound of Formula (I) is selected from a compound of the formula:



32. The compound of claim 1 wherein the compounds are effective antagonists of an integrin receptor.

33. The compound of claim 32 wherein the compound is a selective antagonist of an α_4 integrin receptor.

32
~~33~~ 34. The compound of claim 33 wherein the $\alpha 4$ integrin receptor is selected from the group consisting of the $\alpha 4\beta 1$ and $\alpha 4\beta 7$ integrin receptor.

5 34 35. The compound of claim 32 wherein the compound is an antagonist of at least two $\alpha 4$ integrin receptors.

10 35 36. The compound of claim 35 wherein the two $\alpha 4$ integrin receptors are selected from the group consisting of the $\alpha 4\beta 1$ and $\alpha 4\beta 7$ integrin receptor.

15 37. The compound of claim 1 wherein the compounds are effective agents for the treatment of an integrin mediated disorder ameliorated by selective inhibition of the $\alpha 4\beta 1$ integrin receptor.

20 38. The compound of claim 1 wherein the compounds are effective agents for the treatment of an integrin mediated disorder ameliorated by selective inhibition of the $\alpha 4\beta 7$ integrin receptor.

25 39. The compound of claim 1 wherein the compounds are effective agents for the treatment of an integrin mediated disorder ameliorated by inhibition of the $\alpha 4\beta 1$ and $\alpha 4\beta 7$ integrin receptor.

30 40. The compound of claim 1 wherein the compounds are effective agents for the treatment of integrin mediated disorder selected from the group consisting of inflammatory disorders, autoimmune disorders and cell-proliferative disorders.

41. The compound of claim 40 wherein the integrin mediated disorder is selected from the group consisting of inflammation disorders, autoimmunity disorders, asthma, bronchoconstriction, restenosis, atherosclerosis, psoriasis, rheumatoid arthritis, inflammatory bowel

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disease, irritable bowel disease, irritable bowel syndrome, transplant rejection and multiple sclerosis.

5 42. The compound of claim 40 wherein the integrin mediated disorder is selected from the group consisting of asthma, bronchoconstriction, restenosis, atherosclerosis, psoriasis, rheumatoid arthritis, inflammatory bowel disease, irritable bowel disease, irritable bowel syndrome, transplant rejection and multiple sclerosis.

10 43. The compound of claim 40 wherein the integrin mediated disorder is selected from the group consisting of asthma, bronchoconstriction, restenosis, atherosclerosis, irritable bowel syndrome and multiple sclerosis.

15 36 44. A pharmaceutical composition comprising a compound of claim 1 and a pharmaceutically acceptable carrier.

20 38 45. A pharmaceutical composition made by mixing a compound of claim 1 and a pharmaceutically acceptable carrier.

25 38 46. A method for the treatment of an integrin mediated disorder ameliorated by inhibition of an $\alpha 4$ integrin receptor comprising administering to a subject in need thereof a therapeutically effective amount of a compound of claim 1.

30 47. The method of claim 46 wherein the compound inhibiting the $\alpha 4$ integrin receptor is selected from the group consisting of a selective antagonist of an $\alpha 4$ integrin receptor and an antagonist of at least two $\alpha 4$ integrin receptors.

30 48. The method of claim 47 wherein the $\alpha 4$ integrin receptor is selected from the group consisting of the $\alpha 4\beta 1$ and $\alpha 4\beta 7$ integrin receptor.

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The method of claim 46 wherein the compound inhibiting the $\alpha 4$ integrin receptor is selected from the group consisting of a selective antagonist of the $\alpha 4 \beta 1$ integrin receptor, a selective antagonist of the $\alpha 4 \beta 7$ integrin receptor and an antagonist of the $\alpha 4 \beta 1$ and $\alpha 4 \beta 7$ integrin receptors.

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Said AS

50. The method of claim 46 wherein the integrin mediated disorder is selected from the group consisting of inflammatory disorders, autoimmune disorders and cell-proliferative disorders.

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51. The method of claim 46 wherein the integrin mediated disorder is selected from the group consisting of inflammation disorders, autoimmunity disorders, asthma, bronchoconstriction, restenosis, atherosclerosis, psoriasis, rheumatoid arthritis, inflammatory bowel disease, irritable bowel disease, irritable bowel syndrome, transplant rejection and multiple sclerosis.

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52. The compound of claim 46 wherein the integrin mediated disorder is selected from the group consisting of asthma, bronchoconstriction, restenosis, atherosclerosis, psoriasis, rheumatoid arthritis, inflammatory bowel disease, irritable bowel disease, irritable bowel syndrome, transplant rejection and multiple sclerosis.

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53. The compound of claim 46 wherein the integrin mediated disorder is selected from the group consisting of asthma, bronchoconstriction, restenosis, atherosclerosis, irritable bowel syndrome and multiple sclerosis.

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54.

The method of claim 46 wherein the therapeutically effective amount of the compound of claim 1 is from about 0.01 mg/kg/day to about 300 mg/kg/day.

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Said AS

55.

The method of claim 46 further comprising administering to a subject in need thereof a therapeutically effective amount of the pharmaceutical

composition of claim 44.

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56. The method of claim 55 wherein the therapeutically effective amount of the pharmaceutical composition of claim 44 is from about 0.01 mg/kg/day to about 300 mg/kg/day.

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The compound of claim 1 wherein R₇ is selected from the group consisting tolyl, phenyl and thienyl.

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when R₃ and R₄ comprise a bond and C₁₋₈alkyl or optionally when both R₃ and R₄ are C₁₋₈alkyl, R₃ and R₄ together with the atoms to which each is attached will form a five to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R₃ and R₅ comprise a bond and C₁₋₈alkyl or optionally when both R₃ and R₅ are C₁₋₈alkyl, R₃ and R₅ together with the atoms to which each is attached will form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R₄ and R₅ comprise a bond and C₁₋₈alkyl, or optionally when both R₄ and R₅ are C₁₋₈alkyl, R₄ and R₅ together with the atoms to which each is attached will form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

R₆ is optionally present and is one to three substituents independently selected from the group consisting of halogen, C₁₋₈alkoxy, R₁₀, R₁₂, -N(R₁₁)C(O)-R₁₀, -N(R₁₁)C(O)-R₁₂, -N(R₁₁)SO₂-R₁₀, -N(R₁₁)SO₂-R₁₂, -N(R₁₁)C(O)-N(R₁₁,R₁₀), -N(R₁₁)C(O)-N(R₁₁,R₁₂), -N(R₁₁)C(O)-N(R₁₂,R₁₇), -C(O)-N(R₁₁,R₁₀), -C(O)-N(R₁₁,R₁₂), -C(O)-N(R₁₂,R₁₇), -OC(O)-N(R₁₁,R₁₀), -OC(O)-N(R₁₁,R₁₂), -OC(O)-N(R₁₂,R₁₇), -OC(O)-R₁₀, -OC(O)-R₁₂, -O-R₁₀ and R₁₀-(C₁₋₈)alkoxy;

R₇, R₉, R₁₀ and R₁₄ are independently selected from the group consisting of cycloalkyl, heterocyclyl, aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, C₁₋₈alkylcarbonyl, C₁₋₈alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃; wherein cycloalkyl and heterocyclyl

are optionally substituted with one to three oxo substituents; and, wherein the aryl and heteroaryl substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently selected from the group consisting of halogen, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃;

R₈, R₁₂, R₁₃ and R₁₇ are independently selected from the group consisting of C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, and (halo)₁₋₃(C₁₋₈)alkyl; wherein C₁₋₈alkyl, C₂₋₈alkenyl and C₂₋₈alkynyl are optionally substituted on a terminal carbon with one to three substituents independently selected from R₁₄;

R₁₁ is selected from the group consisting of hydrogen and C₁₋₈alkyl;

A is C₁₋₄alkylene optionally substituted with one to two substituents independently selected from R₁₃;

when R₃ is C₁₋₈alkyl, optionally A and R₃ together with the atoms to which each is attached may form a five to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R₄ is C₁₋₈alkyl, optionally A and R₄ together with the atoms which each is attached may form a five to seven membered monocyclic ring optionally containing one additional heteroatom selected from the group consisting of N, O and S;

when R₅ is C₁₋₈alkyl, optionally A and R₅ together with the atoms which each is attached may form a three to seven membered monocyclic ring optionally containing one to two heteroatoms independently selected from the group consisting of N, O and S; and,

B₁ and B₂ are independently selected from the group consisting of C₁₋₈alkylene

one to three substituents independently selected from R_9 ; provided that R_2 , R_3 , R_4 and R_5 can only be a bond when forming a monocyclic ring wherein the following monocyclic rings may be formed from R_2 , R_3 , R_4 and R_5 :

when R_2 and R_3 comprise a bond and C_{1-8} alkyl or optionally when both R_2 and R_3 are C_{1-8} alkyl, R_2 and R_3 together with the atoms to which each are attached form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_3 and R_4 comprise a bond and C_{1-8} alkyl or optionally when both R_3 and R_4 are C_{1-8} alkyl, R_3 and R_4 together with the atoms to which each are attached form a five to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_3 and R_5 comprise a bond and C_{1-8} alkyl or optionally when both R_3 and R_5 are C_{1-8} alkyl, R_3 and R_5 together with the atoms to which each are attached form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_4 and R_5 comprise a bond and C_{1-8} alkyl or optionally when both R_4 and R_5 are C_{1-8} alkyl, R_4 and R_5 together with the atoms to which each are attached form a four to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

R_6 is optionally present and is one to three substituents independently selected from the group consisting of halogen, C_{1-8} alkoxy, R_{10} , R_{12} , $-N(R_{11})C(O)-R_{10}$, $-N(R_{11})C(O)-R_{12}$, $-N(R_{11})SO_2-R_{10}$, $-N(R_{11})SO_2-R_{12}$, $-N(R_{11})C(O)-N(R_{11}, R_{10})$, $-N(R_{11})C(O)-N(R_{11}, R_{12})$, $-N(R_{11})C(O)-N(R_{12}, R_{17})$, $-C(O)-N(R_{11}, R_{10})$, $-C(O)-N(R_{11}, R_{12})$, $-C(O)-N(R_{12}, R_{17})$, $-OC(O)-N(R_{11}, R_{10})$, $-OC(O)-N(R_{11}, R_{12})$,

$-\text{OC}(\text{O})-\text{N}(\text{R}_{12}, \text{R}_{17})$, $-\text{OC}(\text{O})-\text{R}_{10}$, $-\text{OC}(\text{O})-\text{R}_{12}$, $-\text{O}-\text{R}_{10}$ and $\text{R}_{10}-(\text{C}_{1-8})\text{alkoxy}$;

R_7 , R_9 , R_{10} and R_{14} are independently selected from the group consisting of cycloalkyl, heterocyclyl, aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, $\text{C}_{1-8}\text{alkyl}$, $\text{C}_{2-8}\text{alkenyl}$, $\text{C}_{2-8}\text{alkynyl}$, $\text{C}_{1-8}\text{alkoxy}$, $\text{C}_{1-8}\text{alkylcarbonyl}$, $\text{C}_{1-8}\text{alkoxycarbonyl}$, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, $\text{N}-(\text{C}_{1-8}\text{alkyl})\text{amino}$, $\text{N,N}-(\text{C}_{1-8}\text{dialkyl})\text{amino}$, $-\text{CF}_3$ and $-\text{OCF}_3$, wherein cycloalkyl and heterocyclyl are optionally substituted with one to three oxo substituents; and, wherein the aryl and heteroaryl substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently selected from the group consisting of halogen, $\text{C}_{1-8}\text{alkyl}$, $\text{C}_{2-8}\text{alkenyl}$, $\text{C}_{2-8}\text{alkynyl}$, $\text{C}_{1-8}\text{alkoxy}$, carboxyl, amino, $\text{N}-(\text{C}_{1-8}\text{alkyl})\text{amino}$, $\text{N,N}-(\text{C}_{1-8}\text{dialkyl})\text{amino}$, $-\text{CF}_3$ and $-\text{OCF}_3$;

R_8 , R_{12} , R_{13} and R_{17} are independently selected from the group consisting of $\text{C}_{1-8}\text{alkyl}$, $\text{C}_{2-8}\text{alkenyl}$, $\text{C}_{2-8}\text{alkynyl}$, and $(\text{halo})_{1-3}(\text{C}_{1-8})\text{alkyl}$; wherein $\text{C}_{1-8}\text{alkyl}$, $\text{C}_{2-8}\text{alkenyl}$ and $\text{C}_{2-8}\text{alkynyl}$ are optionally substituted on a terminal carbon with one to three substituents independently selected from R_{14} ;

R_{11} is selected from the group consisting of hydrogen and $\text{C}_{1-8}\text{alkyl}$;

A is $\text{C}_{1-4}\text{alkylene}$ optionally substituted with one to two substituents independently selected from R_{13} ;

when R_3 is $\text{C}_{1-8}\text{alkyl}$, optionally A and R_3 together with the atoms to which each is attached form a five to seven membered monocyclic ring optionally containing one to two additional heteroatoms independently selected from the group consisting of N, O and S;

when R_4 is $\text{C}_{1-8}\text{alkyl}$, optionally A and R_4 together with the atoms to which each is attached form a five to seven membered monocyclic ring optionally

containing one additional heteroatom selected from the group consisting of N, O and S;

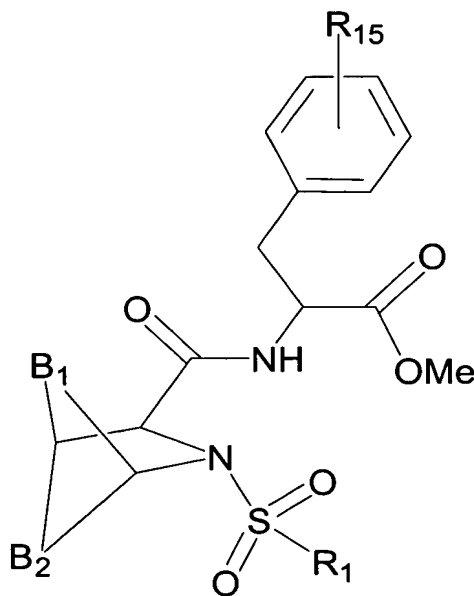
when R_5 is C_{1-8} alkyl, optionally A and R_3 together with the atoms to which each is attached form a three to seven membered monocyclic ring optionally containing one to two heteroatoms independently selected from the group consisting of N, O and S;

B is selected from the group consisting of C_{1-8} alkylene and C_{2-8} alkenylene optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C_{1-8})alkyl, hydroxy(C_{1-8})alkoxy, C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, C_{1-8} alkoxy, carboxyl, amino, N -(C_{1-8} alkyl)amino, N,N -(C_{1-8} dialkyl)amino, $-CF_3$ and $-OCF_3$; and,

n is an integer from 1 to 2;

and pharmaceutically acceptable salts, racemic mixtures, diastereomers and enantiomers thereof.

26. A process for preparing a compound of Formula (III):



Formula (III)

wherein

R_1 is selected from the group consisting of R_7 and R_8 ;

R_7 , R_{10} , and R_{14} are independently selected from the group consisting of cycloalkyl, heterocyclyl, aryl and heteroaryl optionally substituted with one to five substituents independently selected from the group consisting of halogen, C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, C_{1-8} alkoxy, C_{1-8} alkylcarbonyl, C_{1-8} alkoxycarbonyl, carboxyl, aryl, heteroaryl, arylcarbonyl, heteroarylcarbonyl, arylsulfonyl, amino, N -(C_{1-8} alkyl)amino, N,N -(C_{1-8} dialkyl)amino, $-CF_3$ and $-OCF_3$; wherein cycloalkyl and heterocyclyl are optionally substituted with one to three oxo substituents; and, wherein the aryl and heteroaryl substituents and the aryl portion of the arylcarbonyl substituent are optionally substituted with one to five substituents independently selected from the group consisting of halogen, C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, C_{1-8} alkoxy, carboxyl, amino, N -(C_{1-8} alkyl)amino, N,N -(C_{1-8} dialkyl)amino, $-CF_3$ and $-OCF_3$;

R_8 , R_{12} and R_{17} are independently selected from the group consisting of C_{1-8} alkyl, C_{2-8} alkenyl, C_{2-8} alkynyl, and (halo) $_{1-3}$ (C_{1-8} alkyl); wherein C_{1-8} alkyl, C_{2-8} alkenyl and C_{2-8} alkynyl are optionally substituted on a terminal carbon with one to three substituents independently selected from R_{14} ;

R_{150} is selected from the group consisting of hydroxy, amino, NO_2 and R_6 ;

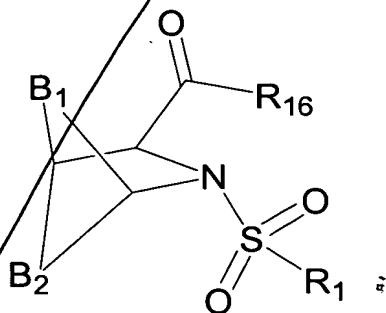
R_6 is optionally present and is one to three substituents independently selected from the group consisting of halogen, C_{1-8} alkoxy, R_{10} , R_{12} , $-N(R_{11})C(O)-R_{10}$, $-N(R_{11})C(O)-R_{12}$, $-N(R_{11})SO_2-R_{10}$, $-N(R_{11})SO_2-R_{12}$, $-N(R_{11})C(O)-N(R_{11}, R_{10})$, $-N(R_{11})C(O)-N(R_{11}, R_{12})$, $-N(R_{11})C(O)-N(R_{12}, R_{17})$, $-C(O)-N(R_{11}, R_{10})$, $-C(O)-N(R_{12}, R_{17})$, $-C(O)-N(R_{11}, R_{12})$, $-OC(O)-N(R_{11}, R_{10})$, $-OC(O)-N(R_{11}, R_{12})$, $-OC(O)-N(R_{12}, R_{17})$, $-OC(O)-R_{10}$, $-OC(O)-R_{12}$, $-O-R_{10}$ and $R_{10}-(C_{1-8})alkoxy$;

R_{11} is selected from the group consisting of hydrogen and C_{1-8} alkyl; and,

B₁ and B₂ are independently selected from the group consisting of C₁₋₈alkylene and C₂₋₈alkenylene optionally substituted with one to two substituents independently selected from the group consisting of halogen, hydroxy, hydroxy(C₁₋₈)alkyl, hydroxy(C₁₋₈)alkoxy, C₁₋₈alkyl, C₂₋₈alkenyl, C₂₋₈alkynyl, C₁₋₈alkoxy, carboxyl, amino, N-(C₁₋₈alkyl)amino, N,N-(C₁₋₈dialkyl)amino, -CF₃ and -OCF₃;

and pharmaceutically acceptable salts, racemic mixtures, diastereomers and enantiomers thereof;

comprising reacting a compound of Formula (IV)

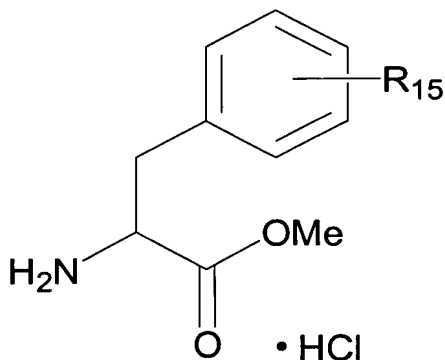


Formula (IV)

wherein

R₁₆ is selected from the group consisting of halogen, mixed anhydride and hydroxy;

with a compound of Formula (V)



Formula (V);